Filed: December 30, 1999

Group Art Unit: 1642

This listing of claims will replace all prior versions, and listings, of claims in the application:

**LISTING OF CLAIMS:** 

Claims 1-10 (canceled)

Claim 11 (previously presented): The method according to claim 51 or 63, wherein the

cancer is a tumor in which a myc gene is amplified.

Claim 12 (previously presented): The method according to claim 51 or 63, wherein the

cancer is a neuroblastoma.

Claims 13-15 (canceled)

Claim 16 (currently amended): The method according to claim 15 51 or 63, wherein the

methylation of CASP8 genomic DNA is detected by methylation polymerase chain reaction

(PCR) assay.

Claims 17-28 (canceled)

Claim 29 (currently amended): A kit for detecting inactivation of a CASP8 gene

expression, comprising CASP8 gene-specific oligonucleotide primers for amplification of at least

a part of the 5' untranslated region of CASP8 genomic DNA, wherein said primers are used in a

methylation polymerase chain reaction (PCR) assay.

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Claims 30-50 (canceled)

Claim 51 (currently amended): A method for diagnosis of a cancer comprising detecting

inactivation of a CASP8 gene expression in a cell from a subject, wherein said inactivation of a

CASP8 gene expression in the cell is indicative of the presence of a cancer and wherein said

method comprises at least one assay selected from the group consisting of detecting a

methylation of CASP8 genomic DNA, detecting the absence of expression of a CASP8 protein,

and detecting the absence of a CASP8 mRNA.

Claims 52-55 (canceled)

Claim 56 (previously presented): A method for detecting inactivation of a CASP8 gene

expression in a primary cancer cell, comprising detecting a methylation of CASP8 genomic

DNA.

Claim 57 (previously presented): The method according to claim 56, wherein the

methylation of CASP8 genomic DNA is detected by methylation polymerase chain reaction

(PCR) assay.

Claim 58 (currently amended): The kit of claim 29, wherein the kit comprises <u>CASP8</u>

gene-specific oligonucleotide PCR primers for amplification of SEQ ID NO: 1 or SEQ ID NO: 2.

Claim 59 (previously presented): The kit of claim 58, wherein the kit comprises at least

one oligonucleotide PCR primer selected from the group consisting of SEQ ID NO: 29, SEQ ID

NO: 30, SEQ ID NO: 31, SEQ ID NO: 32, SEQ ID NO: 33, and SEQ ID NO: 34.

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Claim 60 (previously presented): The method according to claim 56, wherein the

methylation occurs in the 5' untranslated region of CASP8 genomic DNA.

Claim 61 (previously presented): The method according to claim 60, wherein the

methylation occurs in sequences selected from the group consisting of SEQ ID NO: 1 and SEQ

ID NO: 2.

Claim 62 (previously presented): The method according to claim 57, wherein the PCR

assay utilizes at least one of the primer sequences selected from the group consisting of SEQ ID

NO: 29, SEQ ID NO: 30, SEQ ID NO: 31, SEQ ID NO: 32, SEQ ID NO: 33, and SEQ ID NO:

34.

Claim 63 (currently amended): A method for prognosis of a cancer comprising detecting

inactivation of a CASP8 gene expression in a cancer cell from a subject, wherein said

inactivation of a CASP8 gene expression in the cancer cell is indicative of the inefficiency of

apoptosis induced by activated death receptors, chemotherapeutic drugs, or irradiation, and

wherein said method comprises at least one assay selected from the group consisting of detecting

a methylation of CASP8 genomic DNA, detecting the absence of expression of a CASP8 protein,

and detecting the absence of a CASP8 mRNA.

Claims 64-65 (canceled)

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For Examiner's convenience, presented below are pending claims as amended arranged in

order from independent to dependent:

Claim 56 (previously presented): A method for detecting inactivation of a CASP8 gene

expression in a primary cancer cell, comprising detecting a methylation of CASP8 genomic

DNA.

Claim 60 (previously presented): The method according to claim 56, wherein the

methylation occurs in the 5' untranslated region of CASP8 genomic DNA.

Claim 61 (previously presented): The method according to claim 60, wherein the

methylation occurs in sequences selected from the group consisting of SEQ ID NO: 1 and SEQ

ID NO: 2.

Claim 57 (previously presented): The method according to claim 56, wherein the

methylation of CASP8 genomic DNA is detected by methylation polymerase chain reaction

(PCR) assay.

Claim 62 (previously presented): The method according to claim 57, wherein the PCR

assay utilizes at least one of the primer sequences selected from the group consisting of SEO ID

NO: 29, SEQ ID NO: 30, SEQ ID NO: 31, SEQ ID NO: 32, SEQ ID NO: 33, and SEQ ID NO:

34.

Claim 51 (currently amended): A method for diagnosis of a cancer comprising detecting

inactivation of a CASP8 gene expression in a cell from a subject, wherein said inactivation of a

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CASP8 gene expression in the cell is indicative of the presence of a cancer and wherein said method comprises at least one assay selected from the group consisting of detecting a methylation of CASP8 genomic DNA, detecting the absence of expression of a CASP8 protein, and detecting the absence of a CASP8 mRNA.

Claim 63 (currently amended): A method for prognosis of a cancer comprising detecting inactivation of a CASP8 gene expression in a cancer cell from a subject, wherein said inactivation of a CASP8 gene expression in the cancer cell is indicative of the inefficiency of apoptosis induced by activated death receptors, chemotherapeutic drugs, or irradiation, and wherein said method comprises at least one assay selected from the group consisting of detecting a methylation of CASP8 genomic DNA, detecting the absence of expression of a CASP8 protein, and detecting the absence of a CASP8 mRNA.

Claim 11 (previously presented): The method according to claim 51 or 63, wherein the cancer is a tumor in which a myc gene is amplified.

Claim 12 (previously presented): The method according to claim 51 or 63, wherein the cancer is a neuroblastoma.

Claim 16 (currently amended): The method according to claim 15 51 or 63, wherein the methylation of CASP8 genomic DNA is detected by methylation polymerase chain reaction (PCR) assay.

Claim 29 (currently amended): A kit for detecting inactivation of a CASP8 gene expression, comprising <u>CASP8</u> gene-specific oligonucleotide primers for amplification of at least

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a part of the 5' untranslated region of CASP8 genomic DNA, wherein said primers are used in a

methylation polymerase chain reaction (PCR) assay.

Claim 58 (currently amended): The kit of claim 29, wherein the kit comprises <u>CASP8</u>

gene-specific oligonucleotide PCR primers for amplification of SEQ ID NO: 1 or SEQ ID NO: 2.

Claim 59 (previously presented): The kit of claim 58, wherein the kit comprises at least

one oligonucleotide PCR primer selected from the group consisting of SEQ ID NO: 29, SEQ ID

NO: 30, SEQ ID NO: 31, SEQ ID NO: 32, SEQ ID NO: 33, and SEQ ID NO: 34.